Amendments to the Claims

What is claimed is:

1. (Canceled)

2. (Canceled)

3. (Canceled)

4. (Currently Amended) A compound of the Formula I:

or stereoisomers, pharmaceutically acceptable salts, solvates and hydrates thereof, wherein:

- R1 is selected from the group consisting of hydrogen, C₁-C₈ alkyl, C₁-C₈ alkenyl, phenyl, phenyl, and C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl, and wherein C₁-C₈ alkyl is optionally substituted with from one to three substituents independently selected from R1'; and further wherein C₁-C₈ alkenyl, phenyl, and C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl, are each optionally substituted with from one to three substituents independently selected from R2;
- (b) R1' are each independently selected from the group consisting of hydroxy, cyano, nitro, halo, oxo, C₁-C₆ alkyl-COOR12, C₁-C₆ alkoxy, C₁-C₆ haloalkyl, C₁-C₆ haloalkyloxy, C₃-C₇ cycloalkyl, aryloxy, aryl-C₁₋₄-alkyl, C(O)R13, COOR14, OC(O)R15, OS(O)₂R16, N(R17)₂, NR18C(O)R19, NR20SO₂R21, SR22, S(O)R23, S(O)₂R24, and S(O)₂N(R25)₂; R12, R13, R14, R15, R16, R17, R18, R19, R20, R21, R22, R23, R24 and R25 are each independently selected from the group consisting of hydrogen, C₁-C₆ alkyl and aryl;
- (c) R2, R26, R27, R28, and R31 are each independently selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C₁-C₆ alkyl, C₁-C₆ alkyl-COOR12, C₁-C₆ alkoxy, C₁-C₆ haloalkyl, C₁-C₆ haloalkyloxy, C₃-C₇ cycloalkyl, aryloxy, aryl-C₀₋₄-alkyl,

heteroaryl, heterocycloalkyl, C(O)R13, COOR14, OC(O)R15, OS(O)₂R16, N(R17)₂, NR18C(O)R19, NR20SO₂R21, SR22, S(O)R23, S(O)₂R24, and S(O)₂N(R25)₂;

- (d) X is O;
- (e) U is an aliphatic linker;
- (f) Y is selected from the group consisting of C, O, S, NH and a single bond;
- (g) E is C(R3)(R4)A or A and wherein
 - (i) A is selected from the group consisting of carboxyl, , C₁-C₆ alkylnitrile, carboxamide, sulfonamide and acylsulfonamide; wherein sulfonamide, and acylsulfonamide are each optionally substituted with from one to two groups independently selected from R⁷;
 - (ii) each R⁷ is independently selected from the group consisting of hydrogen, C₁-C₆ haloalkyl, aryl C₀-C₄ alkyl and C₁-C₆ alkyl;
 - (iii) R3 is selected from the group consisting of hydrogen, C₁-C₅ alkyl, and C₁-C₅ alkoxy; and
 - (iv) R4 is selected from the group consisting of H, C₁-C₅ alkyl, C₁-C₅ alkoxy, aryloxy, C₃-C₆ cycloalkyl, and aryl C₀-C₄ alkyl, and R3 and R4 are optionally combined to form a C₃-C₄ cycloalkyl, and wherein alkyl, alkoxy, cycloalkyl and aryl-alkyl are each optionally substituted with one to three each independently selected from R26; with the proviso that when R1 is C₁-C₈ alkyl, Y is in a para substituted position with relation to X, and X is selected from the group consisting of a bond and O, then R4 is selected from the group consisting of C₁-C₅ alkoxy, aryloxy, and arylC₀-C₄ alkyl;
- (h) R8 is selected from the group consisting of hydrogen, C₁-C₄ alkyl, C₁-C₄ alkylenyl, and halo;
- (i) R9 is selected from the group consisting of hydrogen, C₁-C₄ alkyl, C₁-C₄ alkylenyl, halo, aryl-C₀-C₄ alkyl, , C₁-C₆ allyl, and OR29, and wherein aryl-C₀-C₄ alkyl are each optionally substituted with from one to three independently selected from R27; R29 is selected from the group consisting of hydrogen and C₁-C₄ alkyl;
- (j) R10 is selected from the group consisting of C₃-C₇ cycloalkyl, aryl-C₀₋₄-alkyl, aryl-C₁₋₄-heteroalkyl, heteroaryl-C₀₋₄-alkyl, C3-C6 cycloalkylaryl-C₀₋₂-alkyl, and aryloxy, provided that when the aliphatic linker, U, is C₁-C₃ alkyl substituted with arylC₁-C₄alkyl, then R10 is selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C₁-C₆ alkyl, C₁-C₆ alkyl-COOR12", C₁-C₆ alkoxy, C₁-C₆ haloalkyl, C₁-C₆ haloalkyl, aryl-C₀₋₄-alkyl, aryl-C₁₋₄-heteroalkyl, heteroaryl-C₀₋₄-

alkyl, C3-C6 cycloalkylaryl- C_{0-2} -alkyl, aryloxy, C(O)R13', COOR14', OC(O)R15', OS(O)₂R16', N(R17')₂, NR18'C(O)R19', NR20'SO₂R21', SR22', S(O)R23', S(O)₂R24', and S(O)₂N(R25')₂; and wherein aryl- C_{0-4} -alkyl, aryl- C_{1-4} -heteroalkyl, heteroaryl- C_{0-4} -alkyl, and C3-C6 cycloalkylaryl- C_{0-2} -alkyl are each optionally substituted with from one to three independently selected from R28

- (j)(k) R10, R11 is are each independently-selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C₁-C₆ alkyl, C₁-C₆ alkyl-COOR12'', C₁-C₆ alkoxy, C₁-C₆ haloalkyl, C₁-C₆ haloalkyloxy, C₃-C₇ cycloalkyl, aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, heteroaryl-C₀₋₄-alkyl, C3-C6 cycloalkylaryl-C₀₋₂-alkyl, aryloxy, C(O)R13', COOR14', OC(O)R15', OS(O)₂R16', N(R17')₂, NR18'C(O)R19', NR20'SO₂R21', SR22', S(O)R23', S(O)₂R24', and S(O)₂N(R25')₂; and wherein aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, heteroaryl-C₀₋₄-alkyl, and C3-C6 cycloalkylaryl-C₀₋₂-alkyl are each optionally substituted with from one to three independently selected from R28; and
- (k)(1) R12', R12'', R13', R14', R15', R16', R17', R18', R19', R20', R21', R22', R23', R24', and R25' are each independently selected from the group consisting of hydrogen, C₁-C₆ alkyl and aryl; or the compound of Formula I is selected from the group consisting of 3-{2-Methyl-4-[5-(4-trifluoromethyl-phenyl)-thiophen-2-ylmethoxy]-phenyl}-propionic acid and 3-{2-Methyl-4-[5-(4-trifluoromethyl-phenyl)-thiophen-2-ylmethoxy]-phenyl}-propionic acid.
- 5. (Canceled)
- 6. (Canceled)
- 7. (Currently Amended) A compound as claimed by Claim 4 Claim 4wherein R4 is selected from the group consisting of C₁-C₅ alkoxy, aryloxy, and arylC₀-C₄ alkyl.
- 8. (Previously Presented) A compound as claimed by Claim 4 wherein Y is O.
- 9. (Peviously Presented) A compound as claimed by Claim 7 wherein Y is C.
- 10. (Currently Amended) A compound as claimed by Claim 7Claim7 wherein Y is S.
- 11. (Canceled).
- 12. (Currently Amended) A compound as claimed by Claim 1 wherein A is carboxyl.
- 13. (Previously Presented) A compound as claimed by Claim 4 wherein R1 is H.
- 14. (Previously Presented) A compound as claimed by Claim 13 wherein A is COOH and R1 is H.

15. (Previously Presented) A compound as claimed by Claim 14 wherein R10 is haloalkyl.

- 16. (Currently Amended) A compound as claimed by Claim 4 Claim 11-wherein R10 is CF₃.
- 17. (Previously Presented) A compound as claimed by Claim 14, wherein R10 is haloalkyloxy.
- 18. (Previously Presented) A compound as claimed by Claim 4 wherein R10 and R11 are each independently selected from the group consisting of hydrogen, halo, oxo, C₁-C₆ alkyl, C₁-C₆ alkyl-COOR12", C₁-C₆ alkoxy, C₁-C₆ haloalkyl, and C₁-C₆ haloalkyloxy.
- 19. (Previously Presented) A compound as claimed by Claim 4 wherein R10 is selected from the group consisting of C₃-C₇ cycloalkyl, aryl-C₀₋₄-alkyl, aryl-C₁₋₄-heteroalkyl, heteroaryl-C₀₋₄-alkyl, C3-C6 cycloalkylaryl-C₀₋₂-alkyl, and aryloxy.
- 20. (Previously Presented) A compound as claimed by Claim 4 wherein R8 and R9 are each independently selected from the group consisting of hydrogen and C₁-C₃ alkyl.
- 21. (Currently Amended) A compound as claimed by Claim 4 wherein R3 R3, and R4 are each independently selected from the group consisting of C₁-C₂ alkyl.
- 22. (Currently Amended) A compound as claimed by <u>Claim 4 Claim 11</u>-wherein <u>R3 -R3</u>, and R4 are each independently selected from the group consisting of hydrogen and C₁-C₂ alkyl.
- 23. (Canceled).
- 24. (Currently Amended) A compound as claimed by <u>Claim 4 Claim 11</u>-wherein U is C₁-C₃ alkyl.
- 25. (Original) A compound as claimed by Claim 24 wherein U is saturated.
- 26. (Original) A compound as claimed by Claim24, wherein U is substituted with C₁-C₃ alkyl.
- 27. (Original) A compound as claimed by Claim 24, wherein U is substituted with arylC₁-C₄alkyl.
- 28. (Canceled)
- 29. (Currently Amended) A compound as claimed by Claim 4Claim 11 wherein R1 is phenyl.
- 30. (Currently Amended) A compound as claimed by <u>Claim 4 Claim 11</u>-represented by the following Structural Formula II:

wherein R33 is <u>arylC₁-C₄ alkyl</u>

selected from the group consisting of hydrogen, C₁-C₃ alkyl, and arylC₀-C₄ alkyl.

- 31. (Canceled)
- 32. (Currently Amended) A compound as claimed by Claim 4 Claim 11 represented by the

following Structural Formula III:_

R33 is selected from the group consisting

of hydrogen, C₁-C₃ alkyl, and arylC₀-C₄ alkyl.

33. (Currently Amended) A compound as claimed by <u>Claim 4 Claim 11</u>-represented by the following Structural Formula IV:

34. (Currently Amended) A compound as claimed by Claim 4 Claim 11 wherein the

headpiece of Formula I is:

- 35. (Currently Amended) A compound as claimed by Claim 11-Claim 4 wherein R4 is selected from the group consisting of C₁-C₅ alkyl, C₁-C₅ alkoxy, aryloxy, C₃-C₆ cycloalkyl, and aryl C₀-C₄ alkyl, and wherein alkyl, alkoxy, cycloalkyl and aryl-alkyl are each optionally substituted with one to three each independently selected from R26.
- 36. (Canceled).
- 37. (Withdrawn) A compound as claimed by Claim 6 wherein A is COOH.
- 38. (Withdrawn Currently Amended) A compound as claimed by <u>Claim 4Claim 1</u>, wherein the compound is selected from the group consisting of (2-Methyl-4-{2-[3-methyl-5-(4-trifluoromethyl-phenyl)-thiophen-2-yl]-propylsulfanyl}-phenoxy)-acetic acid, (2-Methyl-4-{2-[3-methyl-5-(4-trifluoromethyl-phenyl)-thiophen-2-yl]-propylsulfanyl}-phenoxy)-acetic acid, <u>and</u> 3-(2-Methyl-4-{2-[3-methyl-5-(4-trifluoromethyl-phenyl)-thiophen-2-yl]-propylsulfanyl}-phenyl)-propionic acid, <u>and</u> (3-{2-[3-Methyl-5-(4-trifluoromethyl-phenyl)-thiophen-2-yl]-propylsulfanyl}-phenyl)-propionic acid, <u>and</u> (3-{2-[3-Methyl-5-(4-trifluoromethyl-phenyl)-thiophen-2-yl]-propoxy}-phenyl) acetic.
- 39. (Currently Amended) A compound as claimed by Claim 1 Claim 4 that is (3-{2-[3-Methyl-5-(4-trifluoromethyl-phenyl)-thiophen-2-yl]-propoxy}-phenyl)-acetic acid.
- 40. (Previously Presented) A compound as claimed by Claim 4 wherein the compound is selected from the group consisting of

Compound	Name
F F S COOH	3-{2-Methyl-4-[5-(4-trifluoromethyl-phenyl)-thiophen-2-ylmethoxy]-phenyl}-propionic acid
	2 (2) 5 1 1 4 52 1 1 5
F F S COOH	3-{2-Methyl-4-[3-phenyl-5- (4-trifluoromethyl-phenyl)- thiophen-2-ylmethoxy]- phenyl}-propionic acid
FFF	3-{4-[3,5-Bis-(4-trifluoromethyl-phenyl)-thiophen-2-ylmethoxy]-2-methyl-phenyl}-propionic acid.
F S COOH	

- 41. (Previously Presented) A compound as claimed by Claim 4 which is 3-{2-Methyl-4-[5-(4-trifluoromethyl-phenyl)-thiophen-2-ylmethoxy]-phenyl}-propionic acid.
- 42. (Currently Amended) A compound as claimed by <u>Claim 4 Claim 11</u> which is the S conformation.
- 43. (Currently Amended) A compound as claimed by Claim 4 Claim 11 which is the R conformation.
- 44. (Currently Amended) A pharmaceutical composition, comprising as an active ingredient, at least one compound as claimed by Claim 4 Claim 11 together with a pharmaceutically acceptable carrier or diluent.
- 45. (Canceled)
- 46. (Currently Amended) A method of treating diabetes mellitus in a mammal, comprising the step of administering to the mammal in need thereof, a therapeutically effective amount of at least one compound of <u>Claim 4 Claim 11</u>.

47. (Currently Amended) A method of treating Metabolic syndrome in a mammal, comprising the step of administering to the mammal in need thereof a therapeutically effective amount of at least one compound of Claim 4 Claim 11.

- 48. (Currently Amended) A method of selectively modulating a PPAR delta receptor comprising administering a compound as claimed by Claim 4 Claim 11 to a mammal in need thereof.
- 49. (Canceled)
- 50. (Currently Amended) A method of treating atherosclerosis in a mammal, comprising the step of administering to the mammal in need thereof a therapeutically effective amount of at least one compound of Claim 4 Claim 11.
- 51. (Canceled)
- 52. (Canceled)
- 53. (Canceled)
- 54. (Canceled)
- 55. (Canceled)
- 56. (Canceled)
- 57. (Currently Amended) A compound as Claimed by Claim 4 Claim 11 for use as a pharmaceutical.
- 58. (Currently Amended) A compound as claimed by Claim 11 Claim 4 wherein the compound is radiolabeled.
- 59. (Canceled)
- 60. (Canceled)